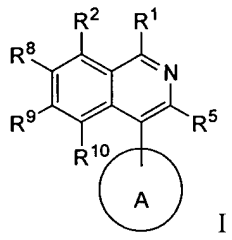
Amendments to the Claims

1. (currently amended) A compound of formula I



I

or a pharmaceutically acceptable salt, ~~crystal form, or hydrate,~~ wherein:

A is

a) an aryl ring selected from phenyl, wherein any stable phenyl ~~aryl~~ ring atom is independently unsubstituted or substituted with

1) halogen,
2) NO_2,
3) CN,
4) $CR^{46}=C(R^{47}R^{48})_2$,
5) $C\equiv C\ R^{46}$,
6) $(CR^iR^j)_rOR^{46}$,
7) $(CR^iR^j)_rN(R^{46}R^{47})$,
8) $(CR^iR^j)_r\ C(O)R^{46}$,
9) $(CR^iR^j)_r\ C(O)OR^{46}$,
10) $(CR^iR^j)_rR^{46}$,
11) $(CR^iR^j)_r\ S(O)_{0-2}R^{61}$,
12) $(CR^iR^j)_r\ S(O)_{0-2}N(R^{46}R^{47})$,
13) $OS(O)_{0-2}R^{61}$,
14) $N(R^{46})C(O)R^{47}$,
15) $N(R^{46})S(O)_{0-2}R^{61}$,
16) $(CR^iR^j)_rN(R^{46})R^{61}$,
17) $(CR^iR^j)_rN(R^{46})R^{61}OR^{47}$,
18) $(CR^iR^j)_rN(R^{46})(CR^kR^l)_sC(O)N(R^{47}R^{48})$,
19) $N(R^{46})(CR^iR^j)_rR^{61}$,
20) $N(R^{46})(CR^iR^j)_rN(R^{47}R^{48})$,
21) $(CR^iR^j)_rC(O)N(R^{47}R^{48})$,

22) oxo,

b) a heteroaryl ring selected from the group consisting of pyridine, pyrimidine, pyrazine, pyridazine, indole, pyrrolopyridine, benzimidazole, benzoxazole, benzothiazole, and benzoxadiazole ~~selected from the group consisting of~~

~~a 5-membered unsaturated monocyclic ring with 1, 2, 3 or 4 heteroatom ring atoms selected from the group consisting or N, O or S,~~

~~a 6-membered unsaturated monocyclic ring with 1, 2, 3 or 4 heteroatom ring atoms selected from the group consisting N, O and S, and~~

~~a 9- or 10-membered unsaturated bicyclic ring with 1, 2, 3 or 4 heteroatom ring atoms selected from the group consisting or N, O or S,~~

wherein any stable S heteroaryl ring atom is unsubstituted or mono- or di-substituted with oxo, and any stable C or N heteroaryl ring atom is independently unsubstituted or substituted with

1) halogen,
2) NO_2,
3) CN,
4) $CR^{46}=C(R^{47}R^{48})_2$,
5) $C\equiv C\ R^{46}$,
6) $(CR^iR^j)_rOR^{46}$,
7) $(CR^iR^j)_rN(R^{46}R^{47})$,
8) $(CR^iR^j)_r\ C(O)R^{46}$,
9) $(CR^iR^j)_r\ C(O)OR^{46}$,
10) $(CR^iR^j)_rR^{46}$,
11) $(CR^iR^j)_r\ S(O)_{0-2}R^{61}$,
12) $(CR^iR^j)_r\ S(O)_{0-2}N(R^{46}R^{47})$,
13) $OS(O)_{0-2}R^{61}$,
14) $N(R^{46})C(O)R^{47}$,
15) $N(R^{46})S(O)_xR^{61}$,
16) $(CR^iR^j)_rN(R^{46})R^{61}$,
17) $(CR^iR^j)_rN(R^{46})R^{61}OR^{47}$,
18) $(CR^iR^j)_rN(R^{46})(CR^kR^l)_sC(O)N(R^{47}R^{48})$,
19) $N(R^{46})(CR^iR^j)_rR^{61}$,
20) $N(R^{46})(CR^iR^j)_rN(R^{47}R^{48})$,
21) $(CR^iR^j)_rC(O)N(R^{47}R^{48})$, or

22) oxo, or

c) a 4-, 5- or 6-membered heterocyclic ring containing 1 or 2 nitrogen atoms, unsubstituted, mono-substituted or di-substituted with C_1-C_6 alkyl;

Y is CH_2, NR^{53}, $NC(O)R^{53}$, $S(O)_{0-2}$ or O;

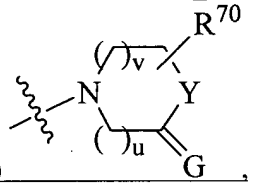
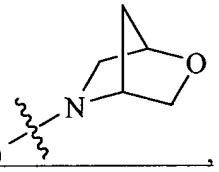
G is H_2 or O;

R^a, R^b, ~~R^e, R^d, R^e, R^f, R^g, R^h~~, R^i, R^j, R^k, and R^l are independently selected from the group consisting of:

1) hydrogen,
2) C_1-C_6 alkyl,
3) halogen,
4) aryl,
5) R^{80},
6) C_3-C_{10} cycloalkyl, and
7) OR^4,

said alkyl, aryl, and cycloalkyl being unsubstituted, monosubstituted with R^7, disubstituted with R^7 and R^{15}, trisubstituted with R^7, R^{15} and R^{16}, or tetrasubstituted with R^7, R^{15}, R^{16} and R^{17};

R^1 is independently selected from:

1) hydrogen,
2) halogen,
3) CN,
4) OR^{40},
5) $N(R^{40}R^{41})$,
6) $C(O)OR^{40}$,
7) R^{81},
8) $S(O)_{0-2}R^6$,
9) $N(R^{40})(CR^aR^b)_nR^6$, wherein $R^6 = R^{83}$,
10) $N(R^{40})(CR^aR^b)_nN(R^{41}R^{42})$,
11)  ,
12)  ,

13) $C(O)N(R^{41}R^{42})$, and

14) a 4-, 5-, or 6-membered heterocyclic ring containing 1 nitrogen atom, unsubstituted, or mono-, di- or tri-substituted with -OH.

~~1) hydrogen,~~

~~2) halogen,~~

~~3) NO_2,~~

~~4) CN,~~

~~5) $CR^{40}=C(R^{41}R^{42})$,~~

~~6) $C\equiv CR^{40}$,~~

~~7) $(CR^aR^b)_nOR^{40}$,~~

~~8) $(CR^aR^b)_nN(R^{40}R^{41})$,~~

~~9) $(CR^aR^b)_nC(O)R^{40}$,~~

~~10) $(CR^aR^b)_nC(O)OR^{40}$,~~

~~11) $(CR^aR^b)_nR^{40}$,~~

~~12) $(CR^aR^b)_nS(O)_{0-2}R^6$,~~

~~13) $(CR^aR^b)_nS(O)_{0-2}N(R^{40}R^{41})$,~~

~~14) $OS(O)_{0-2}R^6$,~~

~~15) $N(R^{40})C(O)R^{41}$,~~

~~16) $N(R^{40})S(O)_{0-2}R^6$,~~

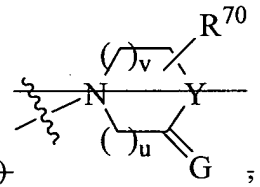
~~17) $(CR^aR^b)_nN(R^{40})R^6$,~~

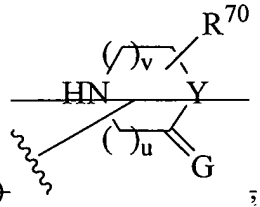
~~18) $(CR^aR^b)_nN(R^{40})R^6OR^{41}$,~~

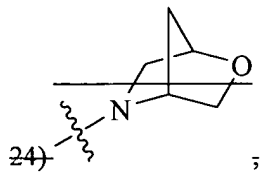
~~19) $(CR^aR^b)_nN(R^{40})(CR^cR^d)_tC(O)N(R^{41}R^{42})$,~~

~~20) $N(R^{40})(CR^aR^b)_nR^6$,~~

~~21) $N(R^{40})(CR^aR^b)_nN(R^{41}R^{42})$,~~

~~22)~~  ;

~~23)~~  ;



~~24)~~ ;

~~25) $(CR^aR^b)_nC(O)N(R^{41}R^{42})$, and~~

~~26) a 4-, 5-, or 6-membered heterocyclic ring containing 1 nitrogen atom, unsubstituted, or mono-, di- or tri-substituted with OH;~~

R^2, R^8, ~~R^9~~ and R^{10} are independently selected from hydrogen and halogen;~~:~~

R^9 is OCH_3 or $OCHF_2$.

~~1) hydrogen,~~

~~2) halogen,~~

~~3) NO_2,~~

~~4) CN,~~

~~5) $CR^{43}=C(R^{44}R^{45})$,~~

~~6) $C\equiv CR^{43}$,~~

~~7) $(CR^eR^f)_pOR^{43}$,~~

~~8) $(CR^eR^f)_pN(R^{43}R^{44})$,~~

~~9) $(CR^eR^f)_pC(O)R^{43}$,~~

~~10) $(CR^eR^f)_pC(O)OR^{43}$,~~

~~11) $(CR^eR^f)_pR^{43}$,~~

~~12) $(CR^eR^f)_pS(O)_{0-2}R^{60}$,~~

~~13) $(CR^eR^f)_pS(O)_{0-2}N(R^{43}R^{44})$,~~

~~14) $OS(O)_{0-2}R^{60}$,~~

~~15) $N(R^{43})C(O)R^{44}$,~~

~~16) $N(R^{43})S(O)_{0-2}R^{60}$,~~

~~17) $(CR^eR^f)_pN(R^{43})R^{60}$,~~

~~18) $(CR^eR^f)_pN(R^{43})R^{60}OR^{44}$,~~

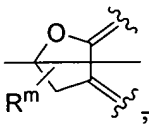
~~19) $(CR^eR^f)_pN(R^{43})(CR^gR^h)_qC(O)N(R^{44}R^{45})$,~~

~~20) $N(R^{43})(CR^eR^f)_pR^{60}$,~~

~~21) $N(R^{43})(CR^eR^f)_pN(R^{44}R^{45})$, and~~

~~22) $(CR^eR^f)_pC(O)N(R^{43}R^{44})$,~~

~~or R^2 and R^8 are independently as defined above, and R^9 and R^{10}, together with the atoms to which they are attached, form the ring~~



~~, where Rm is C1-6alkyl;~~

R4, R40, R41, R42, ~~R43, R44, R45,~~ R46, R47, R48, R49, R50, R51, R52, and R53 are independently selected from:

1) hydrogen,
2) C_1-C_6 alkyl,
3) C_3-C_{10} cycloalkyl,
4) aryl,
5) R^{81},
6) CF_3,
7) C_2-C_6 alkenyl, and
8) C_2-C_6 alkynyl,

said alkyl, aryl, and cycloalkyl is unsubstituted, mono-substituted with R^{18}, di-substituted with R^{18} and R^{19}, tri-substituted with R^{18}, R^{19} and R^{20}, or tetra-substituted with R^{18}, R^{19}, R^{20} and R^{21};

R^5 is independently selected from:

1) hydrogen,
2) halogen,
3) CN,
4) $C(O)N(R^{49}R^{50})$,
5) $C(O)OR^{49}$,
6) $S(O)_{0-2}N(R^{49}R^{50})$,
7) $S(O)_{0-2}R^{62}$,
8) C_1-C_6 alkyl,
9) C_3-C_{10} cycloalkyl,
10) R^{82},

said alkyl, aryl, and cycloalkyl is unsubstituted, mono-substituted with R^{22}, di-substituted with R^{22} and R^{23}, tri-substituted with R^{22}, R^{23} and R^{24}, or tetra-substituted with R^{22}, R^{23}, R^{24} and R^{25};

R6, ~~R60,~~ R61, R62 and R63 are independently selected from:

1) C_1-C_6 alkyl,
2) aryl,
3) R^{83}, and
4) C_3-C_{10} cycloalkyl;

said alkyl, aryl, and cycloalkyl is unsubstituted, mono-substituted with R^{26}, di-substituted with R^{26} and R^{27}, tri-substituted with R^{26}, R^{27} and R^{28}, or tetra-substituted with R^{26}, R^{27}, R^{28} and R^{29};

R^7, R^{15}, R^{16}, R^{17}, R^{18}, R^{19}, R^{20}, R^{21}, R^{22}, R^{23}, R^{24}, R^{25}, R^{26}, R^{27}, R^{28}, R^{29}, and R^{70} are independently selected from:

1) C_1-C_6 alkyl,
2) halogen,
3) OR^{51},
4) CF_3,
5) aryl,
6) C_3-C_{10} cycloalkyl,
7) R^{84},
8) $S(O)_{0-2}N(R^{51}R^{52})$,
9) $C(O)OR^{51}$,
10) $C(O)R^{51}$,
11) CN,
12) $C(O)N(R^{51}R^{52})$,
13) $N(R^{51})C(O)R^{52}$,
14) $S(O)_{0-2}R^{63}$,
15) NO_2, and
16) $N(R^{51}R^{52})$;

R^{80}, R^{81}, R^{82}, R^{83} and R^{84} are independently selected from a group of unsubstituted or substituted heterocyclic rings consisting of a 4-6 membered unsaturated or saturated monocyclic ring with 1, 2, 3 or 4 heteroatom ring atoms selected from the group consisting N, O and S, and a 9- or 10-membered unsaturated or saturated bicyclic ring with 1, 2, 3 or 4 heteroatom ring atoms selected from the group consisting or N, O or S;

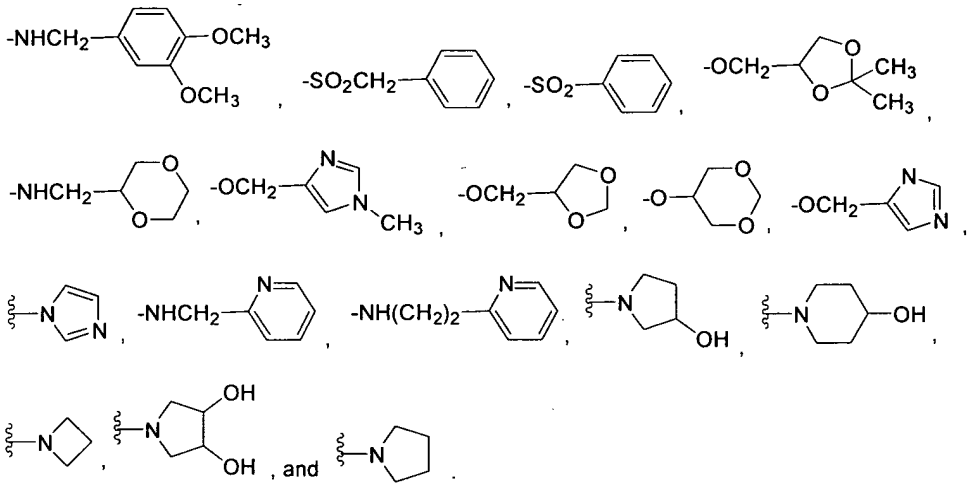
n, ~~p, q,~~ r, s and t are independently 0, 1, 2, 3, 4, 5 or 6;
u is 0, 1 or 2; and
v is 0, 1 or 2.

2.(canceled)·

3. (canceled)·

4. (original) A compound of Claim <u>1</u> ~~3~~, or a pharmaceutically acceptable salt thereof, wherein R^1 is selected from the group consisting of hydrogen, $-SCH_3$, $-SO_2CH_3$, $-NH(CH_2)_3OH$, $-NH(CH_2)_2OH$, $-NH(CH_2)_2OCH_3$,

$-NH(CH_2)_3OCH_3$, $-NH(CH_2)_2NH_2$, $-NH_2$, $-SO_2CH_2CH_3$, -CN, Cl, $-OCH_3$,
$-OCH_2CHCH_2$, $-OCH_2CH(OH)CH_2OH$, $-NHCH_2CHCH_2$, $-CH_3$, $-CH_2CH_2OH$,
$-O(CH_2)_2CHCH_2$, $-O(CH_2)_2CH(OH)(CH_2OH)$, $-NHCH(CH_2OH)_2$,
$-NHCH_2CH(OH)CH_2OH$, $-NH(CH_2)_2CH(OH)CH_2OH$,



5. (original) A compound of Claim 4, or a pharmaceutically acceptable salt thereof, wherein A is selected from the group consisting of

1) phenyl, wherein any stable ring atom is unsubstituted or substituted with halogen,

2) pyridinyl, wherein any stable C ring atom is unsubstituted or substituted with halogen,

3) indolyl, wherein any stable C or N ring atom is unsubstituted or substituted with halogen, and

4) a heterocyclic ring selected from the group consisting of pyrrolidine, piperidine, piperazine, and azetidine, unsubstituted, mono-substituted or di-substituted with C_1-C_6 alkyl.

6. (original) A compound of Claim 5, or a pharmaceutically acceptable salt thereof, wherein R^5 is selected from the group consisting of CN and C_1-C_6 alkyl, wherein said alkyl is unsubstituted, mono-substituted with R^{22}, di-substituted with R^{22} and R^{23}, tri-substituted with R^{22}, R^{23} and R^{24}, or tetra-substituted with R^{22}, R^{23}, R^{24} and R^{25}.

7. (original) A compound of Claim 6, or a pharmaceutically acceptable salt thereof, selected from the group consisting of

[(6-methoxy-4-phenylisoquinolin-3-yl)methyl]dimethylamine,

1-(1-chloro-6-methoxy-4-phenylisoquinolin-3-yl)-N,N-dimethylmethanamine,

{[6-methoxy-1-(methylthio)-4-phenylisoquinolin-3-yl]methyl}dimethylamine,

[6-methoxy-1-(methylsulfonyl)-4-phenylisoquinolin-3-yl]methyl(dimethyl)amine oxide,

1-[6-methoxy-1-(methylsulfonyl)-4-phenylisoquinolin-3-yl]-N,N-dimethylmethanamine,

3-[(dimethylamino)methyl]-6-methoxy-4-phenylisoquinoline-1-carbonitrile,

2,3-Dimethyl-6-methoxy-4-phenylisoquinolinium hydroxide,

6-methoxy-1-(2-methoxyethoxy)-3-methyl-4-phenylisoquinoline,

{3-[(6-methoxy-3-methyl-4-phenylisoquinolin-1-yl)oxy]propyl}amine,

2-[(6-methoxy-3-methyl-4-phenylisoquinolin-1-yl)amino]ethanol,

6-methoxy-3-methyl-1-(methylsulfonyl)-4-phenylisoquinoline,

6-methoxy-N-(2-methoxyethyl)-3-methyl-4-phenylisoquinolin-1-amine,

N-(6-methoxy-3-methyl-4-phenylisoquinolin-1-yl)ethane-1,2-diamine,

6-methoxy-3-methyl-4-phenylisoquinoline,

N-(3,4-dimethoxybenzyl)-6-methoxy-3-methyl-4-phenylisoquinolin-1-amine,

6-methoxy-3-methyl-4-phenylisoquinolin-1-amine,

1-(ethylsulfonyl)-6-methoxy-3-methyl-4-phenylisoquinoline,

1-(benzylsulfonyl)-6-methoxy-3-methyl-4-phenylisoquinoline,

6-methoxy-3-methyl-4-phenyl-1-(phenylsulfonyl)isoquinoline,

6-methoxy-3-methyl-4-phenylisoquinoline-1-carbonitrile,

3-tert-butyl-6-methoxy-1-(2-methoxyethoxy)-4-phenylisoquinoline,

1-chloro-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

6-methoxy-4-phenylisoquinoline-1,3-dicarbonitrile,

1-(allyloxy)-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-(2,3-dihydroxypropoxy)-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

(allylamino)-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

(+/-)-1-[(2,3-dihydroxypropyl)amino]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-{[(2S)-2,3-dihydroxypropyl]amino}-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-{[(2R)-2,3-dihydroxypropyl]amino}-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

(+/-)-1-[(2,2-dimethyl-1,3-dioxolan-4-yl)methoxy]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-{[(4S)-2,2-dimethyl-1,3-dioxolan-4-yl]methoxy}-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-{[(4R)-2,2-dimethyl-1,3-dioxolan-4-yl]methoxy}-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-{[(2R)-2,3-dihydroxypropyl]oxy}-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-{[(2S)-2,3-dihydroxypropyl]oxy}-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

(+/-)-1-{[2,3-dihydroxypropyl]oxy}-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-[(3R)-3-hydroxypyrrolidin-1-yl]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-[(3S)-3-hydroxypyrrolidin-1-yl]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

(+/-)-1-[3-hydroxypyrrolidin-1-yl]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-[cis-3,4-dihydroxypyrrolidin-1-yl]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

6-methoxy-4-phenyl-1-pyrrolidin-1-ylisoquinoline-3-carbonitrile,

6-methoxy-1-(methylsulfonyl)-4-phenylisoquinoline-3-carbonitrile,

6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1,6-dimethoxy-4-phenylisoquinoline-3-carbonitrile,

1-chloro-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

4-(3-fluorophenyl)-6-methoxy-1-methylisoquinoline-3-carbonitrile,

4-(3-fluorophenyl)-1-[(2-hydroxyethyl)amino]-6-methoxyisoquinoline-3-carbonitrile,

1-amino-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

4-(3-fluorophenyl)-1-[(3-hydroxypropyl)amino]-6-methoxyisoquinoline-3-carbonitrile,

1-(but-3-enyloxy)-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

(+/-)-1-(2,3-dihydroxypropoxy)-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(2R)-2,3-dihydroxypropoxy]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(2S)-2,3-dihydroxypropoxy]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

(+/-)-1-(3,4-dihydroxybutoxy)-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

(+/-)-1-[(3R)-3,4-dihydroxybutoxy]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(3S)-3,4-dihydroxybutoxy]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

(+/-)-1-[(1,4-dioxan-2-ylmethyl)amino]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(1,4-dioxan-(2R)-2-ylmethyl)amino]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(1,4-dioxan-(2S)-2-ylmethyl)amino]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

4-(3-fluorophenyl)-6-methoxy-1-[(1-methyl-1H-imidazol-4-yl)methoxy]isoquinoline-3-carbonitrile,

(+/-)-1-(1,3-dioxolan-4-ylmethoxy)-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-(1,3-dioxolan-(4R)-4-ylmethoxy)-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-(1,3-dioxolan-(4S)-4-ylmethoxy)-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-(1,3-dioxan-5-yloxy)-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

4-(3-fluorophenyl)-1-{[2-hydroxy-1-(hydroxymethyl)ethyl]amino}-6-methoxyisoquinoline-3-carbonitrile,

4-(3-fluorophenyl)-1-(1H-imidazol-5-ylmethoxy)-6-methoxyisoquinoline-3-carbonitrile,

1-{[(2R)-2,3-dihydroxypropyl]amino}-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-{[(2S)-2,3-dihydroxypropyl]amino}-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

(+/-)-1-{[2,3-dihydroxypropyl]amino}-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-

carbonitrile,

1-(1H-imidazol-1-yl)-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

6-methoxy-4-phenyl-1-[(pyridin-2-ylmethyl)amino]isoquinoline-3-carbonitrile,

6-methoxy-4-phenyl-1-[(2-pyridin-2-ylethyl)amino]isoquinoline-3-carbonitrile,

(+/-)-1-[(3,4-dihydroxybutyl)amino]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(3R)-(3,4-dihydroxybutyl)amino]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(3S)-(3,4-dihydroxybutyl)amino]-4-(3-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-chloro-4-(2-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

4-(2-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

(+/-)-1-[(2,3-dihydroxypropyl)amino]-4-(2-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(2S)-(2,3-dihydroxypropyl)amino]-4-(2-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

1-[(2R)-(2,3-dihydroxypropyl)amino]-4-(2-fluorophenyl)-6-methoxyisoquinoline-3-carbonitrile,

(+/-)-6-(difluoromethoxy)-1-{[2,3-dihydroxypropyl]amino}-4-(3-fluorophenyl)isoquinoline-3-carbonitrile,

6-(difluoromethoxy)-1-{[(2S)-2,3-dihydroxypropyl]amino}-4-(3-fluorophenyl)isoquinoline-3-carbonitrile,

6-(difluoromethoxy)-1-{[(2R)-2,3-dihydroxypropyl]amino}-4-(3-fluorophenyl)isoquinoline-3-carbonitrile,

(+/-)-6-(difluoromethoxy)-1-{[2,3-dihydroxypropyl]oxy}-4-(3-fluorophenyl)isoquinoline-3-carbonitrile,

6-(difluoromethoxy)-1-{[(2S)-2,3-dihydroxypropyl]oxy}-4-(3-fluorophenyl)isoquinoline-3-carbonitrile,

6-(difluoromethoxy)-1-{[(2R)-2,3-dihydroxypropyl]oxy}-4-(3-fluorophenyl)isoquinoline-3-carbonitrile,

1-(4-hydroxypiperidin-1-yl)-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-azetidin-1-yl-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

(+/-)-1-[trans-3,4-dihydroxypyrrolidin-1-yl]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-[(3R,4R)-3,4-dihydroxypyrrolidin-1-yl]-6-methoxy-4-phenylisoquinoline-3-carbonitrile,

1-[(3S,4S)-3,4-dihydroxypyrrolidin-1-yl]-6-methoxy-4-phenylisoquinoline-3-carbonitrile, and

6-methoxy-N-(3-methoxypropyl)-3-methyl-4-phenylisoquinolin-1-amine.

8. (withdrawn) A method of treating a condition in a mammal, the treatment of which is effected or facilitated by $K_V1.5$ inhibition, which comprises administering a compound of Claim 1 in an amount that is effective at inhibiting $K_V1.5$.

9. (withdrawn) A method of Claim 8, wherein the condition is cardiac arrythmia.

10. (withdrawn) A method of Claim 9, wherein the cardiac arrythmia is atrial fibrillation.

11. (withdrawn) A method of Claim 9, wherein the cardiac arrythmia is selected from the group consisting of atrial flutter, atrial arrhythmia and supraventricular tachycardia.

12. (withdrawn) A method of preventing a condition in a mammal, the prevention of which is effected or facilitated by $K_V1.5$ inhibition, which comprises administering a compound of Claim 1 in an amount that is effective at inhibiting $K_V1.5$.

13. (withdrawn) A method of Claim 12, wherein the condition is cardiac arrythmia.

14. (withdrawn) A method of Claim 13, wherein the cardiac arrythmia is atrial fibrillation.

15. (withdrawn) A method of Claim 13, wherein the cardiac arrythmia is selected from the group consisting of atrial flutter, atrial arrhythmia and supraventricular tachycardia.

16. (withdrawn) A method of Claim 12, wherein the condition is a thromboembolic event.

17. (withdrawn) A method of Claim 16, wherein the thromboembolic event is a stroke.

18. (withdrawn) A method of Claim 12, wherein the condition is congestive heart failure.

19. (currently amended) A pharmaceutical formulation comprising a pharmaceutically acceptable carrier and the compound Claim 1 or a pharmaceutically acceptable salt ~~crystal form or hydrate~~ thereof.

20. (original) A pharmaceutical composition made by combining the compound of Claim 1 and a pharmaceutically acceptable carrier.

21. (withdrawn) A method of treating cardiac arrythmia comprising administering a compound of Claim 1 with a compound selected from one of the classes of compounds consisting of antiarrhythmic agents having Kv1.5 blocking activities, ACE inhibitors, angiotensin II antagonists, cardiac glycosides, L-type calcium channel blockers, T-type calcium channel blockers, selective and nonselective beta blockers, endothelin antagonists, thrombin inhibitors, aspirin, nonselective NSAIDs, warfarin, factor Xa inhibitors, low molecular weight heparin, unfractionated heparin, clopidogrel, ticlopidine, IIb/IIIa receptor antagonists, 5HT receptor antagonists, integrin receptor antagonists, thromboxane receptor antagonists, TAFI inhibitors and P2T receptor antagonists.

22. (withdrawn) A method for inducing a condition of normal sinus rhythm in a patient having atrial fibrillation, which comprises treating the patient with a compound of Claim 1.

23. (withdrawn) A method for treating tachycardia in a patient which comprises treating the patient with an antitachycardia device in combination with a compound of Claim 1.